In re Appln. Of Glorioso et al. Application No. 09/506,301

Amendments to existing claims:

- 1. (Cancelled)
- 2. (Cancelled)
- 3. (Cancelled)
- 4. (Cancelled)
- 5. (Cancelled)
- 6. (Cancelled)
- 7. (Cancelled)
- 8. (Cancelled)
- 9. (Cancelled)
- 10. (Cancelled)
- 11. (Cancelled)
- 12. (Cancelled)
- 13. (Cancelled)
- 14. (Cancelled)
- 15. (Cancelled)
- 16. (Cancelled)
- 21. (Cancelled)
- 22. (Cancelled)
- 23. (Cancelled)
- 24. (Cancelled)
- 25. (Cancelled)
- 26. (Cancelled)
- 27. (Cancelled)
- 28. (Cancelled)
- 29. (Cancelled)
- 30. (Cancelled)
- 31. (New) A recombinant herpes simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adeno-associated virus (AAV) rep polypeptide, wherein the rep polypeptide is conditionally active.
- 32. (New) The recombinant HSV of claim 31, wherein the rep polypeptide is obtained from an AAV rep78, rep68, rep62, or rep40 protein.
- 33. (New) The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep78 protein.

- 34. (New) The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep68 protein.
- 35. (New) The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep62 protein.
- 36. (New) The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep40 protein.
- 37. (New) The recombinant HSV of claim 31, further comprising an Intermediate Terminal Repeat (ITR) cassette, which comprises two AAV-derived ITR sequences flanking a non-ITR polynucleotide.
- 38. (New) The recombinant HSV of claim 37, wherein the rep gene is not within the ITR cassette.
- 39. (New) The recombinant HSV of claim 31, further comprising a cap gene comprising a promoter operatively linked to a polynucleotide sequence encoding an AAV cap polypeptide.
- 40. (New) The recombinant HSV of claim 39, further comprising an ITR cassette, which comprises two AAV-derived ITR sequences flanking a non-ITR polynucleotide.
- 41. (New) The recombinant HSV of claim 40, wherein the rep gene is not within the AAV ITR cassette.
- 42. (New) The recombinant HSV of claim 31, which is deficient for at least one essential HSV.
- 43. (New) The recombinant HSV of claim 42, wherein the essential HSV gene is an immediate early, early or late HSV gene.
- 44. (New) The recombinant HSV of claim 42, wherein the essential HSV gene is ICP27.
- 45. (New) The recombinant HSV of claim 31, wherein the promoter is conditionally active.
- 46. (New) The recombinant HSV of claim 31, wherein the promoter is a tissue specific promoter.
- 47. (New) The recombinant HSV of claim 31, wherein the promoter is an HSV promoter.
- 48. (New) The recombinant HSV of claim 31, which is replication incompetent in cells other than packaging cells.
 - 49. (New) A viral stock comprising the recombinant HSV of claim 31.
- 50. (New) A composition comprising the recombinant HSV of claim 31 and a physiologically-acceptable carrier.
 - 51. (New) The composition of claim 50, which further comprises an ITR cassette.

- 52. (New) The composition of claim 51, wherein the ITR cassette is within an HSV vector.
- 53. (New) The composition of claim 50, further comprising a second HSV that comprises an ITR cassette.
- 54. (New) A recombinant herpes simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adeno-associated virus (AAV) rep polypeptide, a cap gene, which comprises a promoter operatively linked to a polynucleotide sequence encoding an AAV cap polypeptide, and an Intermediate Terminal Cassette (ITR) cassette, which comprises two AAV-derived ITR sequences flanking a non-ITR polynucleotide, wherein the rep polypeptide or the promoter is conditionally active.
- 55. (New) The recombinant HSV of claim 54, wherein the rep polypeptide is obtained from an AAV rep78, rep68, rep62, or rep40 protein.
- 56. (New) The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep78 protein.
- 57. (New) The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep68 protein.
- 58. (New) The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep62 protein.
- 59. (New) The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep40 protein.
- 60. (New) The recombinant HSV of claim 54, wherein the rep gene is not within the AAV ITR cassette.
- 61. (New) The recombinant HSV of claim 54, which is deficient for at least one essential HSV gene.
- 62. (New) The recombinant HSV of claim 61, wherein the essential HSV gene is an immediate early, early or late HSV gene.
- 63. (New) The recombinant HSV of claim 61, wherein the essential HSV gene is ICP27.
- 64. (New) The recombinant HSV of claim 54, wherein the promoter is conditionally active.
- 65. (New) The recombinant HSV of claim 54, wherein the promoter is a tissue specific promoter.
- 66. (New) The recombinant HSV of claim 54, wherein the promoter is an HSV promoter.

- 67. (New) The recombinant HSV of claim 54, which is replication incompetent in cells other than packaging cells.
 - 68. (New) A viral stock comprising the recombinant HSV of claim 54.
- 69. (New) A composition comprising the recombinant HSV of claim 54 and a physiologically-acceptable carrier.
 - 70. (New) The composition of claim 69, which further comprises an ITR cassette.
- 71. (New) The composition of claim 70, wherein the ITR cassette is within an HSV vector.
- 72. (New) The composition of claim 69, further comprising a second HSV that comprises an ITR cassette.
- 73. (New) A recombinant herpes simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adeno-associated virus (AAV) rep polypeptide, wherein the rep polypeptide or the promoter is conditionally active, and wherein the promoter is an inducible promoter.
- 74. (New) A recombinant herpex simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adenoassociated virus (AAV) rep polypeptide, wherein the rep polypeptide is active at a first permissive temperature, and inactive at a second nonpermissive temperature.



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:

Glorioso et al.

Art Unit: 1636 Application No. 09/506,301

Examiner: G. Leffers, Jr.

Filed: February 17, 2000

For: ADENO-ASSOCIATED VIRAL

GENE-TRANSFER VECTOR

SYSTEM

PENDING CLAIMS AFTER AMENDMENTS MADE IN RESPONSE TO OFFICE ACTION DATED JUNE 5, 2002

- 31. A recombinant herpes simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adeno-associated virus (AAV) rep polypeptide, wherein the rep polypeptide is conditionally active.
- 32. The recombinant HSV of claim 31, wherein the rep polypeptide is obtained from an AAV rep78, rep68, rep62, or rep40 protein.
- 33. The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep78 protein.
- 34. The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep68 protein.
- 35. The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep62 protein.
- 36. The recombinant HSV of claim 31, wherein the rep polypeptide is an AAV rep40 protein.
- 37. The recombinant HSV of claim 31, further comprising an Intermediate Terminal Repeat (ITR) cassette, which comprises two AAV-derived ITR sequences flanking a non-ITR polynucleotide.
- 38. The recombinant HSV of claim 37, wherein the rep gene is not within the ITR cassette.
- 39. The recombinant HSV of claim 31, further comprising a cap gene comprising a promoter operatively linked to a polynucleotide sequence encoding an AAV cap polypeptide.
- 40. The recombinant HSV of claim 39, further comprising an ITR cassette, which comprises two AAV-derived ITR sequences flanking a non-ITR polynucleotide.

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- 41. The recombinant HSV of claim 40, wherein the rep gene is not within the AAV ITR cassette.
- 42. The recombinant HSV of claim 31, which is deficient for at least one essential HSV.
- 43. The recombinant HSV of claim 42, wherein the essential HSV gene is an immediate early, early or late HSV gene.
 - 44. The recombinant HSV of claim 42, wherein the essential HSV gene is ICP27.
- 45. The recombinant HSV of claim 31, wherein the promoter is conditionally active.
- 46. The recombinant HSV of claim 31, wherein the promoter is a tissue specific promoter.
- 47. The recombinant HSV of claim 31, wherein the promoter is an HSV promoter.
- 48. The recombinant HSV of claim 31, which is replication incompetent in cells other than packaging cells.
 - 49. A viral stock comprising the recombinant HSV of claim 31.
- 50. A composition comprising the recombinant HSV of claim 31 and a physiologically-acceptable carrier.
 - 51. The composition of claim 50, which further comprises an ITR cassette.
- 52. The composition of claim 51, wherein the ITR cassette is within an HSV vector.
- 53. The composition of claim 50, further comprising a second HSV that comprises an ITR cassette.
- 54. A recombinant herpes simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adeno-associated virus (AAV) rep polypeptide, a cap gene, which comprises a promoter operatively linked to a polynucleotide sequence encoding an AAV cap polypeptide, and an Intermediate Terminal Cassette (ITR) cassette, which comprises two AAV-derived ITR sequences flanking a non-ITR polynucleotide, wherein the rep polypeptide or the promoter is conditionally active.
- 55. The recombinant HSV of claim 54, wherein the rep polypeptide is obtained from an AAV rep78, rep68, rep62, or rep40 protein.
- 56. The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep78 protein.
- 57. The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep68 protein.

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- 58. The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep62 protein.
- 59. The recombinant HSV of claim 54, wherein the rep polypeptide is an AAV rep40 protein.
- 60. The recombinant HSV of claim 54, wherein the rep gene is not within the AAV ITR cassette.
- 61. The recombinant HSV of claim 54, which is deficient for at least one essential HSV gene.
- 62. The recombinant HSV of claim 61, wherein the essential HSV gene is an immediate early, early or late HSV gene.
 - 63. The recombinant HSV of claim 61, wherein the essential HSV gene is ICP27.
- 64. The recombinant HSV of claim 54, wherein the promoter is conditionally active.
- 65. The recombinant HSV of claim 54, wherein the promoter is a tissue specific promoter.
- 66. The recombinant HSV of claim 54, wherein the promoter is an HSV promoter.
- 67. The recombinant HSV of claim 54, which is replication incompetent in cells other than packaging cells.
 - 68. A viral stock comprising the recombinant HSV of claim 54.
- 69. A composition comprising the recombinant HSV of claim 54 and a physiologically-acceptable carrier.
 - 70. The composition of claim 69, which further comprises an ITR cassette.
- 71. The composition of claim 70, wherein the ITR cassette is within an HSV vector.
- 72. The composition of claim 69, further comprising a second HSV that comprises an ITR cassette.
- 73. A recombinant herpes simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adeno-associated virus (AAV) rep polypeptide, wherein the rep polypeptide or the promoter is conditionally active, and wherein the promoter is an inducible promoter.
- 74. A recombinant herpex simplex virus (HSV) comprising a rep gene, which comprises a promoter operatively linked to a polynucleotide encoding an adeno-associated virus (AAV) rep polypeptide, wherein the rep polypeptide is active at a first permissive temperature, and inactive at a second nonpermissive temperature.